

Applicants: Nils Ove Gustavsson *et al.*
Serial No.: 10/627,920
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Docket No.: 28069-585-DIV
(Formerly: 003301-072)

Listing of Claims:

This listing of claims will replace all prior versions and listings of claims in the application:

1.-29. (Canceled)

30. (Previously presented) A pharmaceutically acceptable starch for parenteral administration to a mammal, which

- a) has an amylopectin content in excess of 85 percent by weight, in which the molecular weight of said amylopectin has been reduced by shearing to obtain a molecular weight distribution in which at least 80 percent by weight of the starch lies within the range of 100-4000 kDa;
- b) has a purity of at most 50 µg amino acid nitrogen per gram dry weight of starch; and
- c) can be dissolved in a concentration exceeding 25 percent by weight in water.

31. (Previously presented) A pharmaceutically acceptable starch for parenteral administration to a mammal, which

- a) has an amylopectin content in excess of 85 percent by weight, in which the molecular weight of said amylopectin has been reduced by shearing to obtain a molecular weight distribution in which at least 80 percent by weight of the starch lies within the range of 100-4000 kDa;
- b) has a purity of at most 50 µg amino acid nitrogen per gram dry weight of starch; and
- c) lacks hydroxyethylation.

32. (Previously presented) The starch according to claim 30 or claim 31, which exhibits the ability to gel *in vitro*.

33. (Previously presented) The starch according to claim 31, which exhibits the ability to form microparticles in an emulsion system comprising a two-phase aqueous system.

34. (Previously presented) The starch according to claim 31, which has an endotoxin content of less than 35 EU/g and contains fewer than 100 microorganisms per gram.

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35. (Canceled)
36. (Previously presented) The starch according to claim 31, in which the molecular weight of the amylopectin is within the range of 200-1000 kDa.
37. (Previously presented) The starch according to claim 31, which can be dissolved in water in a concentration exceeding 30% by weight.
38. (Previously presented) The starch according to claim 31, which remains in solution at a temperature of at most 60°C, for a period sufficiently long to allow combining with a substance that is temperature sensitive and/or unstable in organic solvents.
39. (Canceled)
40. (Previously presented) The starch according to claim 31, which, when dissolved in water, solidifies at a temperature of 1-55°C.
41. (Presently amended) The starch according to claim 31, which solidifies when exposed to an initial temperature of 1-10°C, and to a subsequent temperature range of 20-40 °C selected from 20-25°C, or 25-40°C.
42. (Previously presented) Microparticles based on starch as a carrier for a biologically active substance for parenteral administration to a mammal, in which said starch is the starch as defined in claim 31.
43. (Previously presented) Microparticles according to claim 42, which have a mean particle diameter in the range of 10-200 µm.
44. (Previously presented) Microparticles according to claim 42, which exhibit the ability to be dissolved by enzymatic action *in vitro* or eliminated from biological tissue *in vivo*.
45. (Previously presented) Microparticles based on starch as a carrier for a biologically active substance which is a protein for parenteral administration to a mammal, in which said starch is the starch as defined in claim 31.

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46. (Canceled)

47. (Previously presented) A starch having a purity of at most 50 µg amino acid nitrogen per gram dry weight of starch and an endotoxin content of less than 25 EU/g and containing fewer than 100 microorganisms per gram, said starch being pharmaceutically acceptable for injection into a human being and obtainable by a process starting from starch in solid form with an amylopectin content in excess of 85 percent by weight expressed as dry weight of starch comprising the following steps:

- (a) subjecting said solid starch to one or more washings under conditions such that material comprising proteins, lipids and endotoxins surface-localized on the starch, as well as more sparingly soluble proteins, are dissolved while the starch remains undissolved, and separating the starch from the dissolved material, said one or more washings comprising a washing with an aqueous alkaline solution for dissolving water-soluble proteins, lipids and endotoxins and a washing with an aqueous solvent with the ability to dissolve zein for dissolving said more sparingly soluble proteins;
- (b) causing the washed starch obtained from step (a) to dissolve in an aqueous medium;
- (c) subjecting the starch solution to a molecular weight reduction by shearing such that a molecular weight distribution is obtained in which at least 80 percent by weight of the material lies within the range of 100-4000 kDa; and
- (d) removing residual water-soluble proteins from the starch by subjecting the starch solution to ion exchange chromatography, said ion exchange chromatography being performed either before or after the shearing step (c), wherein the starch is pharmaceutically acceptable for injection into a human being.

48. (Previously presented) The starch according to claim 30 or claim 31, wherein parenteral administration is by injection.

49. (Previously presented) The starch according to claim 30 or claim 31, wherein the mammal is a human.

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50. (Previously presented) The starch according to claim 30 or claim 31, wherein, in b), the starch has a purity of at most 20 µg amino acid nitrogen per gram dry weight of starch.
51. (Previously presented) The starch according to claim 30 or claim 31, wherein, in b), the starch has a purity of at most 10 µg amino acid nitrogen per gram dry weight of starch.
52. (Previously presented) The starch according to claim 30 or claim 31, wherein, in b), the starch has a purity of at most 5 µg amino acid nitrogen per gram dry weight of starch.
53. (Previously presented) The starch according to claim 30, in which, in a), the molecular weight of the amylopectin is within the range of 200-1000 kDa.
54. (Previously presented) The starch according to claim 31, in which, in a), the molecular weight of the amylopectin is within the range of 300-600 kDa.
55. (Previously presented) The starch according to claim 31, which can be dissolved in water in a concentration exceeding 40% by weight.
56. (Previously presented) The starch according to claim 31, which can be dissolved in water in a concentration exceeding 45% by weight.
57. (Previously presented) The starch according to claim 31, which remains in solution at a temperature of at most 20-45°C, for a period sufficiently long to allow combining with a substance that is temperature sensitive and/or unstable in organic solvents.
58. (Previously presented) The starch according to claim 31, which remains in solution at a temperature of at most 30-37°C, for a period sufficiently long to allow combining with a substance that is temperature sensitive and/or unstable in organic solvents.
59. (Presently amended) The starch according to any one of claims ~~31~~, 57, or 58, in which the substance that is temperature sensitive and/or unstable in organic solvents is a protein.
60. (Previously presented) The starch according to claim 40, which, when dissolved in water, solidifies at a temperature of 4-37°C.

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61. (Previously presented) The starch according to claim 41, which solidifies when exposed to an initial temperature of 4°C, and to a subsequent temperature of 37°C.
62. (Previously presented) The microparticles according to claim 42, which are parenterally administered by injection.
63. (Previously presented) The microparticles according to claim 43, which have a mean particle diameter in the range of 20-100 µm.
64. (Previously presented) The microparticles according to claim 43, which have a mean particle diameter in the range of 20-80 µm.
65. (New) The starch according to any one of claims 38, 57, or 58, wherein said combining is performed under conditions which are able to retain the bioactivity of the substance.